FILE 'HOME' ENTERED AT 12:23:49 ON 27 AUG 2007

=> file registry

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

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STRUCTURE FILE UPDATES: 26 AUG 2007 HIGHEST RN 945604-45-5 DICTIONARY FILE UPDATES: 26 AUG 2007 HIGHEST RN 945604-45-5

New CAS Information Use Policies, enter HELP USAGETERMS for details.

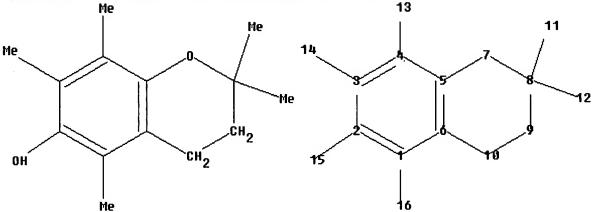
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# http://www.cas.org/support/stngen/stndoc/properties.html

Uploading C:\Program Files\Stnexp\Queries\10789835\_NEW.str



chain nodes :

11 12 13 14 15 16

ring nodes :

1 2 3 4 5 6 7 8 9 10

chain bonds :

1-16 2-15 3-14 4-13 8-11 8-12

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6 5-7 6-10 7-8 8-9 9-10

exact/norm bonds :

2-15 5-7 6-10 7-8 8-9 9-10

exact bonds :

1-16 3-14 4-13 8-11 8-12

normalized bonds :

1-2 1-6 2-3 3-4 4-5 5-6

## Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS

## L1 STRUCTURE UPLOADED

=> d l1

L1 HAS NO ANSWERS

L1 STR

Structure attributes must be viewed using STN Express query preparation.

### => s l1 exa full

FULL SEARCH INITIATED 12:24:28 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 68 TO ITERATE

100.0% PROCESSED 68 ITERATIONS 8 ANSWERS

SEARCH TIME: 00.00.01

L2 8 SEA EXA FUL L1

=> file medline, caplus, wpids, uspatfull

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST 58.25 58.46

FILE 'MEDLINE' ENTERED AT 12:24:41 ON 27 AUG 2007

FILE 'CAPLUS' ENTERED AT 12:24:41 ON 27 AUG 2007
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PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
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FILE 'WPIDS' ENTERED AT 12:24:41 ON 27 AUG 2007 COPYRIGHT (C) 2007 THE THOMSON CORPORATION

FILE 'USPATFULL' ENTERED AT 12:24:41 ON 27 AUG 2007
CA INDEXING COPYRIGHT (C) 2007 AMERICAN CHEMICAL SOCIETY (ACS)

=> s 12

SAMPLE SEARCH INITIATED 12:24:45 FILE 'WPIDS'

SAMPLE SCREEN SEARCH COMPLETED - 62 TO ITERATE

100.0% PROCESSED 62 ITERATIONS 0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*

BATCH \*\*COMPLETE\*\*

PROJECTED ITERATIONS: 384 TO 856
PROJECTED ANSWERS: 0 TO 0

L3 511 L2

=> s 13 and "prostate cancer"

2 FILES SEARCHED...

L4 11 L3 AND "PROSTATE CANCER"

=> d 14 1-11 ibib, abs, hitstr

L4 ANSWER 1 OF 11 MEDLINE on STN

ACCESSION NUMBER: 2003400986 MEDLINE Full-text

DOCUMENT NUMBER: PubMed ID: 12939470

TITLE: Androgen antagonist activity by the antioxidant moiety of

vitamin E, 2,2,5,7,8-pentamethyl-6-chromanol in human

prostate carcinoma cells.

AUTHOR: Thompson Todd A; Wilding George

CORPORATE SOURCE: University of Wisconsin Comprehensive Cancer Center,

University of Wisconsin-Madison, Madison, Wisconsin 53792,

USA.

SOURCE: Molecular cancer therapeutics, (2003 Aug) Vol. 2, No. 8,

pp. 797-803.

Journal code: 101132535. ISSN: 1535-7163.

PUB. COUNTRY:

United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

(RESEARCH SUPPORT, NON-U.S. GOV'T)

(RESEARCH SUPPORT, U.S. GOV'T, NON-P.H.S.)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200406

ENTRY DATE: Entered STN: 27 Aug 2003

Last Updated on STN: 24 Jun 2004 Entered Medline: 21 Jun 2004

AB Antioxidants, such as vitamin E, are being investigated for efficacy in prostate cancer prevention. In this study, we show that the antioxidant moiety of vitamin E, 2,2,5,7,8-pentamethyl-6-chromanol (PMCol), has antiandrogen activity in prostate carcinoma cells. In the presence of PMCol, the androgen-stimulated biphasic growth curve of LNCaP human prostate carcinoma cells was shifted to the right. The PMCol-induced growth shift was similar to that produced by treatment with the pure antiandrogen bicalutamide (i.e., Casodex), indicative of androgen receptor (AR) antagonist activity. The concentration of PMCol used was below the concentration required to affect cell growth or viability in the absence of androgen. Using an AR binding competition assay, PMCol was found to be a potent antiandrogen in both LNCaP

and LAPC4 cells, with an IC(50) of approximately 10 micro M against 1 nM R1881 (methyltrienolone; a stable, synthetic androgen). Prostate-specific antigen release from LNCaP cells produced by androgen exposure with either 0.05 or 1.0 nM R1881 was inhibited 100% and 80%, respectively, by 30 micro M PMCol. Also, PMCol inhibited androgen-induced promoter activation in both LNCaP and LAPC4 cells. However, PMCol did not affect AR protein levels, suggesting that the inhibitory effects of PMCol on androgenic pathways were not due to decreased expression of the AR. Therefore, growth modulation by the antioxidant moiety of vitamin E in androgen-sensitive prostate carcinoma cells is due, at least in part, to its potent antiandrogenic activity.

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L4 ANSWER 2 OF 11 CAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 2005:120717 CAPLUS Full-text
```

DOCUMENT NUMBER: 142:170094

TITLE: Chroman-derived antiandrogens for treatment of

androgen-mediated disorders

INVENTOR(S): Thompson, Todd A.; Wilding, George

PATENT ASSIGNEE(S): Wisconsin Alumni Research Foundation, USA

SOURCE: PCT Int. Appl., 69 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

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KIND DATE APPLICATION NO. DATE
    PATENT NO.
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                                                                _____
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                              -----
    WO 2005011658
                       A2
                                        WO 2004-US5872
                              20050210
                                                               20040227
                       A2 20050210
A3 20050519
    WO 2005011658
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
            CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
            LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
            NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
            TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
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            BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
            ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK,
            TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
    AU 2004260631
                               20050210 AU 2004-260631 20040227
                        A1
                        Al
                               20050210
                                         CA 2004-2517390
                                                                20040227
    CA 2517390
                       A1 20050901 US 2004-789835
A2 20051123 EP 2004-785845
                                                                20040227
    US 2005192342
                                                                20040227
    EP 1596857
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
                                          US 2003-450510P P 20030227
WO 2004-US5872 A 20040227
PRIORITY APPLN. INFO.:
```

## OTHER SOURCE(S): MARPAT 142:170094

- AB Methods for the prevention and/or alleviation of androgen-mediated disorders treatable by administering a chroman-derived antiandrogen compound are provided by the invention. The invention further provides pharmaceutical and nutraceutical compns. containing chroman-derived antiandrogen compds. useful in the prevention and/or alleviation of androgen-mediated disorders, particularly prostate cancer. Compds. of the invention include e.g. 2,2,5,7,8-pentamethyl-6-chromanol.
- IT 950-99-2, 2,2,5,7,8-Pentamethyl-6-chromanol
   RL: ADV (Adverse effect, including toxicity); DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(chroman-derived antiandrogens for treatment of androgen-mediated disorders)

RN 950-99-2 CAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

L4 ANSWER 3 OF 11 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2003:665773 CAPLUS Full-text

DOCUMENT NUMBER:

140:52950

TITLE:

Androgen Antagonist Activity by the Antioxidant Moiety

of Vitamin E, 2,2,5,7,8-Pentamethyl-6-chromanol in

Human Prostate Carcinoma Cells

AUTHOR(S):

Thompson, Todd A.; Wilding, George

CORPORATE SOURCE:

University of Wisconsin Comprehensive Cancer Center and University of Wisconsin Department of Medicine, University of Wisconsin-Madison, Madison, WI, 53792,

USA

SOURCE:

Molecular Cancer Therapeutics (2003), 2(8), 797-803

CODEN: MCTOCF; ISSN: 1535-7163

PUBLISHER:

American Association for Cancer Research

DOCUMENT TYPE: LANGUAGE: Journal English

Antioxidants, such as vitamin E, are being investigated for efficacy in AB prostate cancer prevention. In this study, we show that the antioxidant moiety of vitamin E, 2,2,5,7,8-pentamethyl-6-chromanol (PMCol), has antiandrogen activity in prostate carcinoma cells. In the presence of PMCol, the androgen-stimulated biphasic growth curve of LNCaP human prostate carcinoma cells was shifted to the right. The PMCol-induced growth shift was similar to that produced by treatment with the pure antiandrogen bicalutamide (i.e., Casodex), indicative of androgen receptor (AR) antagonist activity. The concentration of PMCol used was below the concentration required to affect cell growth or viability in the absence of androgen. Using an AR binding competition assay, PMCol was found to be a potent antiandrogen in both LNCaP and LAPC4 cells, with an IC50 of approx. 10 µM against 1 nM R1881 (methyltrienolone; a stable, synthetic androgen). Prostate-specific antigen release from LNCaP cells produced by androgen exposure with either 0.05 or 1.0 nM R1881 was inhibited 100% and 80%, resp., by 30  $\mu M$  PMCol. Also, PMCol inhibited androgen-induced promoter activation in both LNCaP and LAPC4 cells. However, PMCol did not affect AR protein levels, suggesting that the inhibitory effects of PMCol on androgenic pathways were not due to decreased expression of the AR. Therefore, growth modulation by the antioxidant moiety of vitamin E in androgen-sensitive prostate carcinoma cells is due, at least in part, to its potent antiandrogenic activity.

IT 950-99-2, 2,2,5,7,8-Pentamethyl-6-chromanol

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(androgen antagonist activity by the antioxidant moiety of vitamin E, 2,2,5,7,8-pentamethyl-6-chromanol in human prostate carcinoma cells)

RN 950-99-2 CAPLUS

REFERENCE COUNT: 31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 4 OF 11 USPATFULL on STN

ACCESSION NUMBER: 2006:27860 USPATFULL Full-text

TITLE: Novel pathways in the etiology of cancer

INVENTOR(S): Benz, Christopher C., Novato, CA, UNITED STATES
PATENT ASSIGNEE(S): Buck Institute for Age Research (U.S. corporation)

	NUMBER	KIND	DATE	
PATENT INFORMATION:	US 2006024691	Al	20060202	
APPLICATION INFO.:	US 2005-90546	A1	20050324	(11)

			NUMBER	DATE		
PRIORITY	INFORMATION:	US	2004-556774P	20040325	(60)	
		US	2004-580534P	20040616	(60)	
		US	2004-629691P	20041119	(60)	

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: QUINE INTELLECTUAL PROPERTY LAW GROUP, P.C., P O BOX

458, ALAMEDA, CA, 94501, US

NUMBER OF CLAIMS: 30 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 19 Drawing Page(s)

LINE COUNT: 2824

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention pertains to the identification of two novel epithelial signaling pathways in ER-positive breast cancer s and the discovery that the cellular biology and (likely also the clinical outcome) of ER-positive breast cancer cells is unexpectedly altered when these signaling pathways are activated. The first pathway pertains to the discovery that NF-kB activation and/or DNA binding is implicated in the etiology of ER-positive breast (and other) cancers. The second pathway involves ligand-independent quinine-mediated ER activation by posphorylation (e.g. on SER-118 and SER-167 residues of ER) and nuclear translocation of full-length (67 kDA) ER as well as the phorphorylating activation of a truncated and nuclear-localized ER variant (.about.52 kDa).

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2

(pathways in etiol. of cancer, and treatment methods)

RN 950-99-2 USPATFULL

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

L4 ANSWER 5 OF 11 USPATFULL on STN

ACCESSION NUMBER: 2005:221605 USPATFULL Full-text

TITLE: Chroman-derived anti-androgens for treatment of

androgen-mediated disorders

INVENTOR(S): Thompson, Todd A., Madison, WI, UNITED STATES

Wilding, George, Verona, WI, UNITED STATES

PATENT ASSIGNEE(S): Wisconsin Alumni Research Foundation (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 2005192342 A1 20050901

APPLICATION INFO.: US 2004-789835 A1 20040227 (10)

NUMBER DATE

PRIORITY INFORMATION: US 2003-450510P 20030227 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: GODFREY & KAHN, S.C., 780 N. WATER STREET, MILWAUKEE,

WI, 53202, US

NUMBER OF CLAIMS: 24 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 8 Drawing Page(s)

LINE COUNT: 1654

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Methods for the prevention and/or alleviation of androgen-mediated disorders treatable by administering a chroman-derived anti-androgen compound are provided by the present invention. The invention further provides pharmaceutical and nutraceutical compositions containing chroman-derived anti-androgen compounds useful in the prevention and/or alleviation of androgen-mediated disorders, particularly prostate cancer.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2, 2,2,5,7,8-Pentamethyl-6-chromanol

(chroman-derived antiandrogens for treatment of androgen-mediated disorders)

RN 950-99-2 USPATFULL

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

ANSWER 6 OF 11 USPATFULL on STN

ACCESSION NUMBER: 2004:300069 USPATFULL Full-text

Tocopherols, tocotrienols, other chroman and side chain TITLE:

derivatives and uses thereof

Sanders, Bob G., Austin, TX, UNITED STATES INVENTOR(S):

> Kline, Kimberly, Austin, TX, UNITED STATES Hurley, Laurence, Austin, TX, UNITED STATES Gardner, Robb, Austin, TX, UNITED STATES Menchaca, Marla, Austin, TX, UNITED STATES

Yu, Weiping, Austin, TX, UNITED STATES

Ramanan, Puthucode N., Austin, TX, UNITED STATES

Liu, Shenquan, Austin, TX, UNITED STATES Israel, Karen, Austin, TX, UNITED STATES

Research Development Foundation (U.S. corporation) PATENT ASSIGNEE(S):

> NUMBER KIND DATE

-----20041125 PATENT INFORMATION:

US 2003-644418 AJ APPLICATION INFO.: 20030820 (10)

Division of Ser. No. US 2000-502592, filed on 11 Feb RELATED APPLN. INFO.:

2000, GRANTED, Pat. No. US 6770672 Continuation-in-part

of Ser. No. US 1999-404001, filed on 23 Sep 1999,

GRANTED, Pat. No. US 6417223

DATE NUMBER

------PRIORITY INFORMATION: US 1998-101542P 19980923 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

Benjamin Aaron Adler, ADLER & ASSOCIATES, 8011 Candle LEGAL REPRESENTATIVE:

Lane, Houston, TX, 77071

30 NUMBER OF CLAIMS: 1

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 21 Drawing Page(s)

2556 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides an antiproliferative compound having the AB

structural formula ##STR1##

wherein X is oxygen, nitrogen or sulfur; Y is selected from the group consisting of oxygen, nitrogen and sulfur wherein when Y is oxygen or nitrogen, n is 1 and when Y is sulfur, n is 0. Also provided is a method for inducing apoptosis in a cell comprising administering a composition comprising a compound.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2

(preparation of tocopherols, tocotrienols, other chroman and side chain derivs. for use as antitumor agents and for inducing cell apoptosis)

950-99-2 USPATFULL RN

2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME) CN

L4 ANSWER 7 OF 11 USPATFULL on STN

ACCESSION NUMBER: 2004:192666 USPATFULL Full-text

TITLE: Tocopherols, tocotrienols, other chroman and side chain

derivatives and uses thereof

INVENTOR(S): Sanders, Bob G., Austin, TX, United States

Kline, Kimberly, Austin, TX, United States Hurley, Laurence, Austin, TX, United States Gardner, Robb, Austin, TX, United States Menchaca, Marla, Austin, TX, United States Yu, Weiping, Austin, TX, United States

Ramanan, Puthucode N., Austin, TX, United States

Liu, Shenquan, Austin, TX, United States Israel, Karen, Austin, TX, United States

PATENT ASSIGNEE(S): Research Development Foundation, Carson City, NV,

United States (U.S. corporation)

PATENT INFORMATION: APPLICATION INFO.:

US 6770672 B1 20040803 US 2000-502592 20000211 (9)

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 1999-404001, filed on 23 Sep 1999, now patented, Pat. No. US 6417223,

issued on 9 Jul 2002

NUMBER DATE

PRIORITY INFORMATION: US 1998-101543P 19980923 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Fonda, Kathleen K. ASSISTANT EXAMINER: Maier, Leigh C.

LEGAL REPRESENTATIVE: Adler, Benjamin Aaron

NUMBER OF CLAIMS: 4
EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 18 Drawing Figure(s); 21 Drawing Page(s)

LINE COUNT: 2359

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention provides an antiproliferative compound having the

structural formula ##STR1##

wherein X is oxygen, nitrogen or sulfur; Y is selected from the group consisting of oxygen, nitrogen and sulfur wherein when Y is oxygen or nitrogen, n is 1 and when Y is sulfur, n is 0. Also provided is a method for inducing apoptosis in a cell comprising administering a composition comprising a compound.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2, 2,2,5,7,8-Pentamethyl-6-chromanol

(preparation of tocopherols, tocotrienols, other chroman and side chain derivs. for therapeutic use in prevention and treatment of cancer)

950-99-2 USPATFULL RN

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

ANSWER 8 OF 11 USPATFULL on STN

ACCESSION NUMBER:

2004:127448 USPATFULL Full-text

TITLE:

Tocopherols, tocotrienols, other chroman and side chain

derivatives and uses thereof

INVENTOR(S):

Sanders, Bob G., Austin, TX, UNITED STATES Kline, Kimberly, Austin, TX, UNITED STATES

Yu, Weiping, Austin, TX, UNITED STATES

PATENT ASSIGNEE(S):

Research Development Foundation (U.S. corporation)

NUMBER	KIND	DATE

PATENT INFORMATION:

A1 20040520 US 2004097431

APPLICATION INFO.:

(10) US 2003-695275 A1 20031028

Division of Ser. No. US 2001-8066, filed on 5 Nov 2001, RELATED APPLN. INFO.: GRANTED, Pat. No. US 6703384 Continuation-in-part of Ser. No. US 2000-502592, filed on 11 Feb 2000, PENDING Continuation-in-part of Ser. No. US 1999-404001, filed

on 23 Sep 1999, GRANTED, Pat. No. US 6417223

NUMBER DATE \_\_\_\_\_\_

PRIORITY INFORMATION:

US 1998-101542P 19980923 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

Benjamin Aaron Adler, Ph.D., J.D., Adler & Associates,

8011 Candle Lane, Houston, TX, 77071

NUMBER OF CLAIMS:

17

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

11 Drawing Page(s)

LINE COUNT:

2605

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides an antiproliferative compound having a AB structural formula ##STR1##

> where X and Y independently are oxygen, nitrogen or sulfur; R.sup.1 is alkyl, alkenyl, alkynyl, aryl, heteroaryl, carboxylic acid, carbóxylate, carboxamide, ester, thioamide, thiolacid, thiolester, saccharide, alkoxylinked saccharide, amine, sulfonate, sulfate, phosphate, alcohol, ethers or nitrites; R.sup.2 and R.sup.3 are hydrogen or R.sup.4; R.sup.4 is methyl, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzylester, saccharide or amine; and R.sup.5 is alkenyl; where when Y is nitrogen, said nitrogen is substituted with R.sup.6, wherein R.sup.6 is hydrogen or methyl. Also provided are methods for treating a cell proliferative disease and for

inducing apoptosis in a cell comprising administering this compound is also provided.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2

(preparation of tocopherols, tocotrienols, other chroman and side chain derivs. for use as antitumor agents and for inducing cell apoptosis)

RN 950-99-2 USPATFULL

2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME) CN

ANSWER 9 OF 11 USPATFULL on STN

ACCESSION NUMBER: 2002:280579 USPATFULL Full-text

Tocopherols, tocotrienols, other chroman and side chain TITLE:

derivatives and uses thereof

Sanders, Bob G., Austin, TX, UNITED STATES INVENTOR(S):

Kline, Kimberly, Austin, TX, UNITED STATES Hurley, Laurence, Austin, TX, UNITED STATES Gardner, Robb, Austin, TX, UNITED STATES Menchaca, Marla, Austin, TX, UNITED STATES

Yu, Weiping, Austin, TX, UNITED STATES

Ramanan, Puthucode N., Austin, TX, UNITED STATES

Liu, Shenquan, Austin, TX, UNITED STATES Israel, Karen, Austin, TX, UNITED STATES

Research Development Foundation (U.S. corporation) PATENT ASSIGNEE(S):

NUMBER KIND DATE -----PATENT INFORMATION: US 2002156024 A1 20021024 B2 US 6645998 20031111 A1 APPLICATION INFO.: US 2002-122019 20020412 (10)

Division of Ser. No. US 1999-404001, filed on 23 Sep RELATED APPLN. INFO.:

1999, GRANTED, Pat. No. US 6417223

DATE NUMBER

US 1998-101542P 19980923 (60) PRIORITY INFORMATION:

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

Benjamin Aaron Adler, ADLER & ASSOCIATES, 8011 Candle LEGAL REPRESENTATIVE:

Lane, Houston, TX, 77071

NUMBER OF CLAIMS: 20 EXEMPLARY CLAIM:

14 Drawing Page(s) NUMBER OF DRAWINGS:

LINE COUNT: 2170

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides an antiproliferative compound having the

structural formula ##STR1##

wherein X is oxygen, nitrogen or sulfur; R.sup.1 is alkyl, alkenyl, alkynyl, aryl, heteroaryl, carboxylic acid, carboxylate, carboxamide, ester, thioamide, thiolacid, thiolester, saccharide, alkoxy-linked saccharide, amine, sulfonate, sulfate, phosphate, alcohol, ethers and nitriles; R.sup.2 is hydrogen, methyl, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzylester, saccharide and amine; R.sup.3 is selected from the group consisting of hydrogen, methyl, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzylester, saccharide and amine; R.sup.4 is of methyl, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzylester, saccharide and amine; and R.sup.5 is alkyl, alkenyl, alkynyl, aryl, heteroaryl, carboxyl, amide and ester. Also provided is a method for inducing apoptosis in a cell comprising administering a composition comprising a compound.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2

RN

(preparation of tocopherols, tocotrienols, other chroman and side chain derivs. for use as antitumor agents and for inducing cell apoptosis)
950-99-2 USPATFULL

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

L4 ANSWER 10 OF 11 USPATFULL on STN

ACCESSION NUMBER: 2002:199098 USPATFULL Full-text

TITLE: Tocopherols, tocotrienols, other chroman and side chain

derivatives and uses thereof

INVENTOR(S): Sanders, Bob G., Austin, TX, UNITED STATES

Kline, Kimberly, Austin, TX, UNITED STATES Yu, Weiping, Austin, TX, UNITED STATES

NUMBER KIND DATE

PATENT INFORMATION: US 2002107207 A1 20020808 US 6703384 B2 20040309

APPLICATION INFO.: US 2001-8066 A1 20011105 (10)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 2000-502592, filed

on 11 Feb 2000, PENDING Continuation-in-part of Ser. No. US 1999-404001, filed on 23 Sep 1999, PENDING

NUMBER DATE

PRIORITY INFORMATION: US 1998-101542P 19980923 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Benjamin Aaron Adler, ADLER & ASSOCIATES, 8011 Candle

Lane, Houston, TX, 77071

NUMBER OF CLAIMS: 24 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 12 Drawing Page(s)

LINE COUNT: 2606

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention provides an antiproliferative compound having a

structural formula ##STR1##

where X and Y independently are oxygen, nitrogen or sulfur; R.sup.1 is alkyl, alkenyl, alkynyl, aryl, heteroaryl, carboxylic acid, carboxylate, carboxamide, ester, thioamide, thiolacid, thiolester, saccharide, alkoxylinked saccharide, amine, sulfonate, sulfate, phosphate, alcohol, ethers or nitrites; R.sup.2 and R.sup.3 are hydrogen or R.sup.4; R.sup.4 is methyl, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzylester, saccharide or amine; and R.sup.5 is alkenyl; where when Y is nitrogen, said nitrogen is substituted with R.sup.6, wherein R.sup.6 is hydrogen or methyl. Also provided are methods for treating a cell proliferative disease and for inducing apoptosis in a cell comprising administering this compound is also provided.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2, 2,2,5,7,8-Pentamethyl-6-chromanol

(preparation of tocopherols, tocotrienols, other chromans and side chain derivs. as potential antiproliferative, proapoptotic agents for the treatment of cancer)

RN 950-99-2 USPATFULL

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

L4 ANSWER 11 OF 11 USPATFULL on STN

ACCESSION NUMBER: 2002:168253 USPATFULL <u>Full-text</u>

TITLE: Tocopherols, tocotrienols, other chroman and side chain

derivatives and uses therof

INVENTOR(S): Sanders, Bob G., Austin, TX, United States

Kline, Kimberly, Austin, TX, United States Hurley, Laurence, Austin, TX, United States Gardner, Robb, Austin, TX, United States Menchaca, Marla, Austin, TX, United States

Yu, Weiping, Austin, TX, United States Ramanan, Puthucode N., Austin, TX, United States

Liu, Shenquan, Austin, TX, United States Israel, Karen, Austin, TX, United States

PATENT ASSIGNEE(S): Research Development Foundation, Carson City, NV,

United States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 6417223 B1 20020709
APPLICATION INFO.: US 1999-404001 19990923 (9)

NUMBER DATE

PRIORITY INFORMATION: US 1998-101542P 19980923 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Wilson, James O. ASSISTANT EXAMINER: Maier, Leigh C.

LEGAL REPRESENTATIVE: Adler, Benjamin Aaron

NUMBER OF CLAIMS: 3 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 14 Drawing Figure(s); 14 Drawing Page(s)

LINE COUNT: 1959

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention provides an antiproliferative compound having the

structural formula ##STR1##

wherein X is oxygen, nitrogen or sulfur; R.sup.1 is alkyl, alkenyl, alkynyl, aryl, heteroaryl, carboxylic acid, carboxylate, carboxamide, ester, thioamide, thiolacid, thiolester, saccharide, alkoxy-linked saccharide, amine, sulfonate, sulfate, phosphate, alcohol, ethers and nitriles; R.sup.2 is hydrogen, methyl, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzylester, saccharide and amine; R.sup.3 is selected from the group consisting of hydrogen, methyl, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzylester, saccharide and amine; R.sup.4 is of methyl, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzylester, saccharide and amine; and R.sup.5 is alkyl, alkenyl, alkynyl, aryl, heteroaryl, carboxyl, amide and ester. Also provided is a method for inducing apoptosis in a cell comprising administering a composition comprising a compound.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2

(preparation of tocopherols, tocotrienols, other chroman and side chain derivs. for use as antitumor agents and for inducing cell apoptosis)

RN 950-99-2 USPATFULL

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

=> d his

(FILE 'HOME' ENTERED AT 12:23:49 ON 27 AUG 2007)

FILE 'REGISTRY' ENTERED AT 12:24:10 ON 27 AUG 2007

L1 STRUCTURE UPLOADED

L2 8 S L1 EXA FULL

FILE 'MEDLINE, CAPLUS, WPIDS, USPATFULL' ENTERED AT 12:24:41 ON 27 AUG

2007

L3 511 S L2

L4 11 S L3 AND "PROSTATE CANCER"

=> s 13 and "androgen-dependent"

L5 1 L3 AND "ANDROGEN-DEPENDENT"

## => d 15 ibib, abs, hitstr

L5 ANSWER 1 OF 1 USPATFULL on STN

ACCESSION NUMBER: 2005:221605 USPATFULL Full-text

TITLE: Chroman-derived anti-androgens for treatment of

androgen-mediated disorders

INVENTOR(S): Thompson, Todd A., Madison, WI, UNITED STATES

Wilding, George, Verona, WI, UNITED STATES

PATENT ASSIGNEE(S): Wisconsin Alumni Research Foundation (U.S. corporation)

APPLICATION INFO.: US 2004-789835 A1 20040227 (10)

NUMBER DATE

PRIORITY INFORMATION: US 2003-450510P 20030227 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: GODFREY & KAHN, S.C., 780 N. WATER STREET, MILWAUKEE,

WI, 53202, US

NUMBER OF CLAIMS: 24 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 8 Drawing Page(s)

LINE COUNT: 1654

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Methods for the prevention and/or alleviation of androgen-mediated disorders treatable by administering a chroman-derived anti-androgen compound are provided by the present invention. The invention further provides pharmaceutical and nutraceutical compositions containing chroman-derived anti-androgen compounds useful in the prevention and/or alleviation of androgen-mediated disorders, particularly prostate cancer.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2, 2,2,5,7,8-Pentamethyl-6-chromanol

(chroman-derived antiandrogens for treatment of androgen-mediated disorders)

RN 950-99-2 USPATFULL

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

Me Me Me

#### => d his

(FILE 'HOME' ENTERED AT 12:23:49 ON 27 AUG 2007)

FILE 'REGISTRY' ENTERED AT 12:24:10 ON 27 AUG 2007

STRUCTURE UPLOADED L1

8 S L1 EXA FULL L2

> FILE 'MEDLINE, CAPLUS, WPIDS, USPATFULL' ENTERED AT 12:24:41 ON 27 AUG 2007

L3 511 S L2

11 S L3 AND "PROSTATE CANCER" L4

L5 1 S L3 AND "ANDROGEN-DEPENDENT"

## => FIL REGISTRY

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L6 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2007 ACS on STN

RN 950-99-2 REGISTRY

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME) OTHER CA INDEX NAMES:

CN 6-Chromanol, 2,2,5,7,8-pentamethyl- (6CI, 7CI, 8CI) OTHER NAMES:

CN  $\alpha$ -C-1-Chromanol

CN 2,2,5,7,8-Pentamethyl-6-chromanol

CN 2,2,5,7,8-Pentamethyl-6-hydroxychroman

CN 6-Hydroxy-2,2,5,7,8-pentamethylchroman

CN Chroman C1

CN Chromane C1

CN Chromanol

CN NSC 226236

CN PMC

CN TMC 5

MF C14 H20 O2

CI COM

LC STN Files: AGRICOLA, ANABSTR, BEILSTEIN\*, BIOSIS, BIOTECHNO, CA, CAOLD, CAPLUS, CASREACT, CHEMCATS, CHEMINFORMRX, CHEMLIST, CSCHEM, DDFU, DRUGU, EMBASE, MEDLINE, RTECS\*, SPECINFO, TOXCENTER, USPAT2, USPATFULL (\*File contains numerically searchable property data)

DT.CA CAplus document type: Conference; Journal; Patent; Report

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); PRP (Properties); RACT (Reactant or reagent); USES (Uses); NORL (No role in record)

RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological study); FORM (Formation, nonpreparative); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses); NORL (No role in record)

RLD.NP Roles for non-specific derivatives from non-patents: BIOL (Biological study); FORM (Formation, nonpreparative); PREP (Preparation); PROC (Process); PRP (Properties); USES (Uses)

# \*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

405 REFERENCES IN FILE CA (1907 TO DATE)

12 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

405 REFERENCES IN FILE CAPLUS (1907 TO DATE)

14 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

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=> s 16

'RN' IS NOT A VALID FIELD CODE L7 510 L6

=> s 17 and (cancer or tumor)

L8 22 L7 AND (CANCER OR TUMOR)

=> d 18 1-22 ibib, abs, hitstr

L8 ANSWER 1 OF 22 MEDLINE on STN

ACCESSION NUMBER: 2003400986 MEDLINE Full-text

DOCUMENT NUMBER: PubMed ID: 12939470

TITLE: Androgen antagonist activity by the antioxidant moiety of

vitamin E, 2,2,5,7,8-pentamethyl-6-chromanol in human

prostate carcinoma cells.

AUTHOR: Thompson Todd A; Wilding George

CORPORATE SOURCE: University of Wisconsin Comprehensive Cancer Center,

University of Wisconsin-Madison, Madison, Wisconsin 53792,

USA.

SOURCE: Molecular cancer therapeutics, (2003 Aug) Vol. 2, No. 8,

pp. 797-803.

Journal code: 101132535. ISSN: 1535-7163.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE) (RESEARCH SUPPORT, NON-U.S. GOV'T)

(RESEARCH SUPPORT, U.S. GOV'T, NON-P.H.S.)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200406

ENTRY DATE: Entered STN: 27 Aug 2003

Last Updated on STN: 24 Jun 2004

Antioxidants, such as vitamin E, are being investigated for efficacy in AB prostate cancer prevention. In this study, we show that the antioxidant moiety of vitamin E, 2,2,5,7,8-pentamethyl-6-chromanol (PMCol), has antiandrogen activity in prostate carcinoma cells. In the presence of PMCol, the androgen-stimulated biphasic growth curve of LNCaP human prostate carcinoma cells was shifted to the right. The PMCol-induced growth shift was similar to that produced by treatment with the pure antiandrogen bicalutamide (i.e., Casodex), indicative of androgen receptor (AR) antagonist activity. The concentration of PMCol used was below the concentration required to affect cell growth or viability in the absence of androgen. Using an AR binding competition assay, PMCol was found to be a potent antiandrogen in both LNCaP and LAPC4 cells, with an IC(50) of approximately 10 micro M against 1 nM R1881 (methyltrienolone; a stable, synthetic androgen). Prostate-specific antigen release from LNCaP cells produced by androgen exposure with either 0.05 or 1.0 nM R1881 was inhibited 100% and 80%, respectively, by 30 micro M PMCol. Also, PMCol inhibited androgen-induced promoter activation in both LNCaP and LAPC4 cells. However, PMCol did not affect AR protein levels, suggesting that the inhibitory effects of PMCol on androgenic pathways were not due to decreased expression of the AR. Therefore, growth modulation by the antioxidant moiety of vitamin E in androgen-sensitive prostate carcinoma cells is due, at least in part, to its potent antiandrogenic activity.

L8 ANSWER 2 OF 22 MEDLINE on STN

ACCESSION NUMBER: 95232770 MEDLINE Full-text

DOCUMENT NUMBER: PubMed ID: 7716769

TITLE: Phenoxyl radicals of etoposide (VP-16) can directly oxidize

intracellular thiols: protective versus damaging effects of

phenolic antioxidants.

AUTHOR: Tyurina Y Y; Tyurin V A; Yalowich J C; Quinn P J; Claycamp

H G; Schor N F; Pitt B R; Kagan V E

CORPORATE SOURCE: Department of Environmental and Occupational Health,

University of Pittsburgh, Pennsylvania 15238, USA.

SOURCE: Toxicology and applied pharmacology, (1995 Apr) Vol. 131,

No. 2, pp. 277-88.

Journal code: 0416575. ISSN: 0041-008X.

PUB. COUNTRY: United States

DOCUMENT TYPE: (COMPARATIVE STUDY)

Journal; Article; (JOURNAL ARTICLE)
(RESEARCH SUPPORT, NON-U.S. GOV'T)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199505

ENTRY DATE: Entered STN: 24 May 1995

Last Updated on STN: 3 Feb 1997 Entered Medline: 18 May 1995

AB Phenolic compounds can act as radical scavengers due to their ability to donate a mobile hydrogen to peroxyl radicals producing a phenoxyl radical if the phenoxyl radical formed in the radical scavenging reaction efficiently interacts with vitally important biomolecules, then this interaction may result in cytotoxic effects rather than in antioxidant protection. In the present work we have chosen two model compounds—a phenolic antitumor drug, VP-16, known to be highly cytotoxic, and a homolog of vitamin E, 2,2,5,7,8—pentamethyl-6-hydroxychromane (PMC)—as typical representatives of phenoxyl radicals to study interactions of their phenoxyl radicals with intracellular thiols. Using a water—soluble source of peroxyl radicals, the azo-initiator 2,2'-azobis(2- aminodinopropane) (AAPH), we found that both PMC and VP-16 are very efficient scavengers of peroxyl radicals as evidenced by their ability to inhibit AAPH—induced chemiluminescence of luminol and oxidation of PnA

incorporated into DOPC liposomes. Both PMC and VP-16 were also able to protect against AAPH-induced oxidative degradation of DNA in nuclei from human leukemic K562 cells. In contrast, there was a dramatic difference in the ability of VP-16 and PMC to protect GSH against AAPH-induced oxidation: while PMC inhibited AAPH-induced oxidation of GSH in a concentration-dependent manner, VP-16 did not protect GSH against oxidation. We hypothesized that this was due to different reactivities of the phenoxyl radicals formed by AAPH-derived peroxyl radicals from VP-16 and PMC toward GSH. To substantiate this hypothesis, we compared interactions of the phenoxyl radicals generated from VP-16 and PMC with intracellular thiols in K562 cell homogenates. While the PMC phenoxyl radicals were only slightly affected by thiols, the VP-16 phenoxyl radicals were reduced by thiols. This is evidenced by (i) a significant inhibition of the tyrosinase-induced VP-16 consumption upon addition of K562 cell homogenates, (ii) a depletion of endogenous thiols in K562 cell homogenates induced by VP-16+tyrosinase, (iii) a transient disappearance of the VP-16 phenoxyl radical signal from the ESR spectra and its reappearance after depletion of endogenous thiols, and (iv) elimination of the lag period for the appearance of the VP-16 phenoxyl radical ESR signal subsequent to depletion of thiols by mersalyl acid. To evaluate the contribution of GSH and protein thiols to reduction of the VP-GSH-peroxidase + cumeme hydroperoxide to specifically deplete endogenous GSH. (ABSTRACT TRUNCATED AT 400 WORDS)

1

L8 ANSWER 3 OF 22 CAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 2006:101964 CAPLUS Full-text

DOCUMENT NUMBER: 144:184652

TITLE: Novel pathways in the etiology of cancer,

and treatment methods

INVENTOR(S):
Benz, Christopher C.

PATENT ASSIGNEE(S): Buck Institute for Age Research, USA

SOURCE: U.S. Pat. Appl. Publ., 49 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2006024691	A1	20060202	US 2005-90546	20050324
PRIORITY APPLN. INFO.:			US 2004-556774P I	20040325
			US 2004-580534P F	20040616
			US 2004-629691P F	20041119

The invention pertains to the identification of two novel epithelial signaling pathways in ER-pos. breast cancers and the discovery that the cellular biol. and (likely also the clin. outcome) of ER-pos. breast cancer cells is unexpectedly altered when these signaling pathways are activated. The first pathway pertains to the discovery that NF-kB activation and/or DNA binding is implicated in the etiol. of ER-pos. breast (and other) cancers. The second pathway involves ligand-independent quinine-mediated ER activation by phosphorylation (e.g. on SER-118 and SER-167 residues of ER) and nuclear translocation of full-length (67 kDA) ER as well as the phorphorylating activation of a truncated and nuclear-localized ER variant (.apprx.52 kDa). Also disclosed are methods for identifying patients likely to respond to hormonal therapy and for selecting a therapeutic regimen for the treatment of cancer.

IT 950-99-2

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pathways in etiol. of cancer, and treatment methods)

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

RN

L8 ANSWER 4 OF 22 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2005:120717 CAPLUS <u>Full-text</u>

DOCUMENT NUMBER: 142:170094

TITLE: Chroman-derived antiandrogens for treatment of

androgen-mediated disorders

INVENTOR(S): Thompson, Todd A.; Wilding, George

PATENT ASSIGNEE(S): Wisconsin Alumni Research Foundation, USA

SOURCE: PCT Int. Appl., 69 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.					KIN	D :	DATE			APPLICATION NO.					D	ATE			
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	WO	2005	0116	58	A2 20050210 WO 2004-US5872						20	00402	227						
	WO	2005	0116	58		A3		2005	0519										
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			CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,	
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	AU	2004	2606	31		A1		2005	0210		AU 2	004-	2606	31		2	00402	227	
	CA	2517	390			A1		2005	0210		CA 2	004-	2517	390		2	00402	227	
	US	2005	1923	42		A1		2005	0901	1	US 2	004-	78983	35		2	0040	227	
	EP	1596	857			A2		2005	1123	]	EP 2	004-	78584	45		2	0040	227	
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]	PRIORIT	Y APP								US 2003-450510P P 20030227									
										1	WO 2	004-1	JS58'	72	1	A 2	00402	227	

ADDITON MICH

OTHER SOURCE(S): MARPAT 142:170094

AB Methods for the prevention and/or alleviation of androgen-mediated disorders treatable by administering a chroman-derived antiandrogen compound are provided by the invention. The invention further provides pharmaceutical and nutraceutical compns. containing chroman-derived antiandrogen compds. useful in the prevention and/or alleviation of androgen-mediated disorders, particularly prostate cancer. Compds. of the invention include e.g. 2,2,5,7,8-pentamethyl-6-chromanol.

IT 950-99-2, 2,2,5,7,8-Pentamethyl-6-chromanol

RL: ADV (Adverse effect, including toxicity); DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(chroman-derived antiandrogens for treatment of androgen-mediated disorders)

RN 950-99-2 CAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

L8 ANSWER 5 OF 22 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2004:618733 CAPLUS Full-text

DOCUMENT NUMBER:

141:174332

TITLE:

Preparation of tocopherols, tocotrienols, other

chroman and side chain derivatives for therapeutic use

in the prevention and treatment of cancer

INVENTOR(S):

Sanders, Bob G.; Kline, Kimberly; Hurley, Laurence; Gardner, Robb; Menchaca, Marla; Yu, Weiping; Ramanan,

Puthucode N.; Liu, Shenquan; Israel, Karen

PATENT ASSIGNEE(S):

SOURCE:

Research Development Foundation, USA

U.S., 48 pp., Cont.-in-part of U.S. Ser. No. 404,001.

CODEN: USXXAM

DOCUMENT TYPE:

LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

PAT	ENT NO	).			KIN	)	DATE			APPL:	ICAT:	ION	NO.		Di	ATE	
US	 677067	2			B1	-	2004	0803		US 2	000-	 5025	 92		2	0000:	211
US	641722	:3			B1		2002	0709		US 1:	999-	4040	01		1	9990	923
CA	239980	2			Al		2001	0816		CA 2001-2399802 20010209							209
WO	200105	888	39		A1		2001	0816		WO 2	001-1	US41	68		2	0010	209
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CN	152970	1			Α		2004									0010	
RU	226367	2			C2		2005									0010	
US	200210	720	7		A1		2002	8080		US 2	001-	8066			2	0011	105
US	670338	4			B2		2004	0309									

20030820 A1 20041125 US 2003-644418 US 2004235938 20040520 US 2003-695275 20031028 US 2004097431 A1 US 1998-101543P P 19980923 PRIORITY APPLN. INFO.: A2 19990923 US 1999-404001 US 1998-101542P P 19980923 US 2000-502592 Α 20000211 WO 2001-US4168 W 20010209 US 2001-8066 A3 20011105

OTHER SOURCE(S):

MARPAT 141:174332

GI

AB Chroman derivs., such as I [X = O, S, NR6; Y = O, NR6; R1 = carboxyalkyl, carboxyalkenyl, etc.; R2, R3, R4 = H, Me, alkyl, etc.; R5 = alkyl, alkenyl, etc.; R6 = H, alkyl], were prepared for use in antitumor pharmaceutical compns. for inducing apoptosis in a cell, particularly a cancer cell. Thus,  $\alpha$ -tocopherol derivative II was prepared in 88% yield by a reaction of BrCH2CO2Me with (R,R,R)- $\alpha$ -tocopherol using NaOH in DMF. The prepared chromans were assayed for growth inhibitory and apoptotic activity against a variety of human cancer cell lines.

RN 950-99-2 CAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 6 OF 22 CAPLUS COPYRIGHT 2007 ACS ON STN ACCESSION NUMBER: 2003:665773 CAPLUS Full-text DOCUMENT NUMBER: 140:52950

Androgen Antagonist Activity by the Antioxidant Moiety TITLE:

of Vitamin E, 2,2,5,7,8-Pentamethyl-6-chromanol in

Human Prostate Carcinoma Cells

AUTHOR (S):

Thompson, Todd A.; Wilding, George

CORPORATE SOURCE:

University of Wisconsin Comprehensive Cancer Center and University of Wisconsin Department of Medicine, University of Wisconsin-Madison, Madison, WI, 53792,

USA

SOURCE:

Molecular Cancer Therapeutics (2003), 2(8), 797-803

CODEN: MCTOCF; ISSN: 1535-7163

PUBLISHER:

American Association for Cancer Research

DOCUMENT TYPE:

Journal

LANGUAGE:

English

Antioxidants, such as vitamin E, are being investigated for efficacy in prostate cancer prevention. In this study, we show that the antioxidant moiety of vitamin E, 2,2,5,7,8-pentamethyl-6-chromanol (PMCol), has antiandrogen activity in prostate carcinoma cells. In the presence of PMCol, the androgen-stimulated biphasic growth curve of LNCaP human prostate carcinoma cells was shifted to the right. The PMCol-induced growth shift was similar to that produced by treatment with the pure antiandrogen bicalutamide (i.e., Casodex), indicative of androgen receptor (AR) antagonist activity. The concentration of PMCol used was below the concentration required to affect cell growth or viability in the absence of androgen. Using an AR binding competition assay, PMCol was found to be a potent antiandrogen in both LNCaP and LAPC4 cells, with an IC50 of approx. 10 µM against 1 nM R1881 (methyltrienolone; a stable, synthetic androgen). Prostate-specific antigen release from LNCaP cells produced by androgen exposure with either 0.05 or 1.0 nM R1881 was inhibited 100% and 80%, resp., by 30 µM PMCol. Also, PMCol inhibited androgen-induced promoter activation in both LNCaP and LAPC4 cells. However, PMCol did not affect AR protein levels, suggesting that the inhibitory effects of PMCol on androgenic pathways were not due to decreased expression of the AR. Therefore, growth modulation by the antioxidant moiety of vitamin E in androgen-sensitive prostate carcinoma cells is due, at least in part, to its potent antiandrogenic activity.

950-99-2, 2,2,5,7,8-Pentamethyl-6-chromanol IT

> RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(androgen antagonist activity by the antioxidant moiety of vitamin E, 2,2,5,7,8-pentamethyl-6-chromanol in human prostate carcinoma cells)

RN 950-99-2 CAPLUS

2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME) CN

REFERENCE COUNT:

THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS 31 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 7 OF 22 CAPLUS COPYRIGHT 2007 ACS on STN 2002:595501 CAPLUS Full-text ACCESSION NUMBER:

DOCUMENT NUMBER:

TITLE:

Preparation of tocopherols, tocotrienols, other

chromans and side chain derivs. as potential antiproliferative and proapoptotic agents Sanders, Bob G.; Kline, Kimberly; Yu, Weiping

Research Development Foundation, USA

U.S. Pat. Appl. Publ., 44 pp., Cont.-in-part of U.S. SOURCE:

Ser. No. 502,592. CODEN: USXXCO

DOCUMENT TYPE:

INVENTOR(S):

LANGUAGE:

GI

Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT ASSIGNEE(S):

		APPLICATION NO.	DATE					
US 2002107207 US 6703384		US 2001-8066	20011105					
US 6417223		US 1999-404001	19990923					
CN 1706838	A 20051214	CN 2005-10003855	19990923					
US 6770672	B1 20040803	US 2000-502592	20000211					
US 2002156024	A1 20021024	US 2002-122019	20020412					
	B2 20031111	0808 US 2001-8066 20011105 0309 0709 US 1999-404001 19990923 0214 CN 2005-10003855 19990923 0803 US 2000-502592 20000211 024 US 2002-122019 20020412 0515 WO 2002-US35147 20021101 0515 BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM,						
WO 2003039461	A2 20030515	WO 2002-US35147	20021101					
WO 2003039461								
			K, SL, TJ, TM,					
	UG, UZ, VN, YU,	•						
		•						
PRIORITY APPLN. INFO.:	A1 20040520							
PRIORITY APPLIN. INFO.:								
			· · · · · · ·					
		WO 2002-US35147	W 20021101					
OTHER SOURCE(S):	MARPAT 137:14065		<del> </del>					

AB Derivs. of tocopherol, tocotrienol and other chromans of formula I (X and Y independently are oxygen, nitrogen or sulfur; when Y is nitrogen, nitrogen is substituted with R6 and R6 = H or Me; R1 = alkyl, alkenyl, alkynyl, aryl, heteroaryl, carboxylic acid, carboxylate, carboxamide, ester, thioamide, thiolacid, thiol ester, saccharide, alkoxy-linked saccharide, amine, sulfonate, sulfate, phosphate, alc., ethers or nitrites; R2, R3 = hydrogen or R4; R4 = Me, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzyl ester, saccharide or amine; and R5 = alkenyl) were prepared as antiproliferative and proapoptotic agents for the potential treatment of cell proliferative diseases. Thus, α-tocopherol was treated with Me bromoacetate and NaOH in N, N-dimethylformamide to give II. II showed effective growth inhibitory properties (apoptotic inducing) in a wide variety of human cancer cell lines, including breast, prostate, cervical, and ovarian cancers with EC50 values ranging from 1-20 μg/mL.

IT 950-99-2, 2,2,5,7,8-Pentamethyl-6-chromanol

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of tocopherols, tocotrienols, other chromans and side chain derivs. as potential antiproliferative, proapoptotic agents for the treatment of cancer)

RN 950-99-2 CAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

L8 ANSWER 8 OF 22 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2001:597976 CAPLUS Full-text

DOCUMENT NUMBER: 135:166941

TITLE: Preparation of tocopherols, tocotrienols, other

chroman and side chain derivatives that induce cell apoptosis for therapeutic use as antiproliferative

agents

INVENTOR(S): Sanders, Robert G.; Kline, Kimberly; Hurley, Laurence;

Gardner, Robb; Menchaca, Marla; Yu, Weiping; Ramanan,

Puthucode N.; Liu, Shenquan; Israel, Karen

PATENT ASSIGNEE(S): Research Development Foundation, USA

SOURCE: PCT Int. Appl., 120 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

PATENT	NO.			KIN	) ·1	DATE		2	APPL	ICAT:	ION I	NO.		D	ATE	•
WO 2003	.0588	 89		A1	<b>-</b>	2001	0816	,	WO 2	001-1	US41	68		2	 0010:	209
W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,
	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	GM,	HR,
	нп	TD	TT.	TN	TS	JP.	KE	KG	KЪ	KB	K 7.	T.C.	T.K.	T.R	LS.	LT.

LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG US 2000-502592 US 6770672 B1 20040803 20000211 CA 2001-2399802 CA 2399802 A1 20010816 20010209 20010209 EP 2001-909008 EP 1254130 A1 20021106 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR 20040212 JP 2001-558439 20010209 JP 2004504268 Т NZ 2001-520798 20010209 NZ 520798 Α 20040528 20051110 RU 2002-124135 20010209 RU 2263672 C2 US 2000-502592 Α 20000211 PRIORITY APPLN. INFO.: P 19980923 US 1998-101543P US 1999-404001 A2 19990923 WO 2001-US4168 W 20010209

OTHER SOURCE(S):

MARPAT 135:166941

GI

AB Tocopherol analogs, such as I [X = 0, NH, S; Y = 0, NH, S; R1 = alkyl,alkenyl, alkynyl, aryl, heteroaryl, carboxyl, carboxamide, thiocarboxyl, etc.; R2, R3, R4 = H, Me, benzyl, carboxyl, carboxamide, amine, saccharide; R5 = alkyl, alkenyl, alkynyl, aryl, heteroaryl, carboxyl, carboxamide], were prepared for pharmaceutical use as antiproliferative agents which induce cell apoptosis for treatment of cancers and diseases involving cell proliferation, such as autoimmune diseases, psoriasis, etc.. Thus,  $(R,R,R)-\alpha$ -tocopherol derivative II was prepared in 88% yield by condensation of  $(R,R,R)-\alpha$ tocopherol and BrCH2CO2Me in DMF using NaOH followed by hydrolysis with 5 N The prepared tocopherol analogs were tested for their ability to induce apoptosis in a number of cancer cell lines, such as breast, cervical, colon, prostate, etc.

950-99-2 IT

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of tocopherols, tocotrienols, other chromans that induce cell apoptosis for therapeutic use as antiproliferative agents)

RN 950-99-2 CAPLUS

2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME) CN

REFERENCE COUNT: 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 9 OF 22 CAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 2000:209907 CAPLUS Full-text

ACCESSION NUMBER:
DOCUMENT NUMBER:

132:237223

TITLE:

Preparation of tocopherols, tocotrienols, other

chroman and side chain derivatives for use as antitumor agents and for inducing cell apoptosis

INVENTOR(S): Kline, Kimberly; Sanders, Bob G.; Hurley, Laurence; Gardner, Robb; Menchaca, Marla; Yu, Weiping; Ramanan,

Puthucode N.; Liu, Shenquan; Israel, Karen

PATENT ASSIGNEE(S):

Research Development Foundation, USA

SOURCE:

PCT Int. Appl., 101 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

PA	PATENT NO.				KINI	)	DATE			APP:	LICAT	ION I	. 00			DATE	
WO	2000	0167	72		A1		2000	0330	1	WO :	1999-1	US21	778			19990	923
	W:	ΑE,	AL,	AM,	ΑT,	AU,	AZ,	BA,	BB,	ВG	, BR,	BY,	CA,	CH,	CN	, CU,	CZ,
		DE,	DK,	EE,	ES,	FI,	GB,	GD,	GE,	GH	, GM,	HR,	ΗŪ,	ID,	$_{ t IL}$	, IN,	IS,
		JP,	KE,	KG,	KP,	KR,	KZ,	LC,	LK,	LR	, LS,	LT,	LU,	LV,	MD	, MG,	MK,
		MN,	MW,	MX,	NO,	NZ,	PL,	PT,	RO,	RU	, SD,	SE,	SG,	SI,	SK	, SL,	ТJ,
		TM,	TR,	TT,	UA,	ŬĠ,	UΖ,	VN,	YU,	ZA	, ZW						
	RW:	GH,	GM,	KE,	LS,	MW,	SD,	SL,	SZ,	TZ	, UG,	ZW,	AT,	BE,	CH	, CY,	DE,
		DK,	ES,	FI,	FR,	GB,	GR,	ΙE,	IT,	LU	, MC,	NL,	PT,	SE,	BF	, BJ,	CF,
		CG,	CI,	CM,	GΑ,	GN,	GW,	ML,	MR,	NE	, SN,	TD,	TG				
CA	2345	079			A1		2000	0330		CA	1999-	2345	079			19990	923
AU	9961	553			A1		2000	0410		AU	1999-	6155	3			19990	923
AU	7570	13			B2		2003	0130									
EF	1115	398			A1		2001	0718		EP .	1999-	9483	52			19990	923
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR	, IT,	LI,	LU,	NL,	SE	, MC,	PT,
		IE,	SI,	LT,	LV,	FI,	RO										
CN	1325	303			A		2001	1205		CN	1999-	8128	29			19990	923
	2002						2002	0820		JP :	2000-	5737	33			19990	923
NZ	5107	32			Α		2004	0130		NZ	1999-	5107	32			19990	923
RU	2232	758			C2		2004	0720		RU	2001-	1110	19			19990	923
CN	1706						2005	1214		CN	2005-	1000	3855			19990	923
II	1420	82			Α		2005	1218		$_{ m IL}$	1999-	1420	82			19990	923
TW	5926	95			В		2004	0621		TW	1999-	8812	0073			19991	117
ZA	2001	0020	57		Α		2002	0319		ZA	2001-	2057				20010	313
PRIORIT	Y APP	LN.	INFO	. :						US	1998-	1015	42P		P	19980	923
										CN	1999-	8128	29				
										WO	1999-	US21	778	1	W	19990	923

OTHER SOURCE(S): MARPAT 132:237223

GI

$$R^3$$
 $R^4$ 
 $R^5$ 
 $R^1$ 
 $R^2$ 
 $R^5$ 
 $R^6$ 
 $R^1$ 
 $R^2$ 
 $R^2$ 
 $R^2$ 
 $R^3$ 
 $R^4$ 
 $R^5$ 
 $R^6$ 
 $R^6$ 

Chromans I [R1 = alkyl, alkenyl, alkynyl, aryl, herteroaryl, carboxyl, carboxamide, thioamide, saccharide, amine, sulfate, phosphate, etc.; R2, R3, R4 = H, Me, benzylcarboxylate, saccharide, amino, etc.; R5 = alkyl, alkenyl, alkynyl, aryl, herteroaryl, carboxyl, carboxamide; X = O, NH, S] were prepared for pharmaceutical use as antitumor agents and cell apoptosis inducing agents. Thus, tocopherol derivative II (R1 = CH2CO2H, X = O) was prepared in 88% yield via O-alkylation of (+)- $\alpha$ -tocopherol with Me bromoacetate. The prepared chromans were tested for cell apoptosis activity against a variety of cancer cell lines.

IT 950-99-2

RL: RCT (Reactant); RACT (Reactant or reagent)
(preparation of tocopherols, tocotrienols, other chroman and side chain derivs. for use as antitumor agents and for inducing cell apoptosis)

RN 950-99-2 CAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 10 OF 22 CAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 1999:602318 CAPLUS Full-text

DOCUMENT NUMBER:

131:295249

TITLE:

Mechanism-based chemopreventive strategies against

etoposide-induced acute myeloid leukemia: free

radical/antioxidant approach

AUTHOR (S):

Kagan, Valerian E.; Yalowich, Jack C.; Borisenko,
Grigory G.; Tyurina, Yulia Y.; Tyurin, Vladimir A.;

Thampatty, Padmakumari; Fabisiak, James P.

CORPORATE SOURCE:

Departments of Environmental and Occupational Health and Pharmacology and University of Pittsburgh Cancer Institute, University of Pittsburgh, Pittsburgh, PA, USA SOURCE: Molecular Pharmacology (1999), 56(3), 494-506

CODEN: MOPMA3; ISSN: 0026-895X

PUBLISHER: American Society for Pharmacology and Experimental

Therapeutics

DOCUMENT TYPE: Journal LANGUAGE: English

Etoposide (VP-16) is extensively used to treat cancer, yet its efficacy is AB calamitously associated with an increased risk of secondary acute myelogenous leukemia. The mechanisms for the extremely high susceptibility of myeloid stem cells to the leukemogenic effects of etoposide have not been elucidated. We propose a mechanism to account for the etoposide-induced secondary acute myelogenous leukemia and nutritional strategies to prevent this complication of etoposide therapy. We hypothesize that etoposide phenoxyl radicals (etoposide-O) formed from etoposide by myeloperoxidase are responsible for its genotoxic effects in bone marrow progenitor cells, which contain constitutively high myeloperoxidase activity. Here, we used purified human myeloperoxidase, as well as human leukemia HL60 cells with high myeloperoxidase activity and provide evidence of the following. 1. Etoposide undergoes one-electron oxidation to etoposide-O· catalyzed by both purified myeloperoxidase and myeloperoxidase activity in HL60 cells; formation of etoposide-O·radicals is completely blocked by myeloperoxidase inhibitors, cyanide and azide. 2. Intracellular reductants, GSH and protein sulfhydryls (but not phospholipids), are involved in myeloperoxidase-catalyzed etoposide redox-cycling that oxidizes endogenous thiols; pretreatment of HL60 cells with a maleimide thiol reagent, ThioGlol, prevents redox-cycling of etoposide-O. radicals and permits their direct ESR detection in cell homogenates. VP-16 redox-cycling by purified myeloperoxidase (in the presence of GSH) or by myeloperoxidase activity in HL60 cells is accompanied by generation of thiyl radicals, GS., determined by HPLC assay of 5,5-dimethyl-1- pyrroline glytathionyl N-oxide glytathionyl nitrone adducts. 3. Ascorbate directly reduces etoposide-O·, thus competitively inhibiting etoposide-O·-induced thiol oxidation Ascorbate also diminishes etoposide-induced topo II-DNA complex formation in myeloperoxidase-rich HL60 cells (but not in HL60 cells with myeloperoxidase activity depleted by pretreatment with succinyl acetone). A vitamin E homolog, 2,2,5,7,8-pentamethyl-6-hydroxychromane, a hindered phenolic compound whose phenoxyl radicals do not oxidize endogenous thiols, effectively competes with etoposide as a substrate for myeloperoxidase, thus preventing etoposide-O·-induced redox-cycling. We conclude that nutritional antioxidant strategies can be targeted at minimizing etoposide conversion to etoposide-O·, thus minimizing the genotoxic effects of the radicals in bone marrow myelogenous progenitor cells, i.e., chemoprevention of etoposideinduced acute myelogenous leukemia.

IT 950-99-2

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(mechanism-based chemopreventive strategies against etoposide-induced acute myeloid leukemia: free radical/antioxidant approach)

RN 950-99-2 CAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

REFERENCE COUNT: 42 THERE ARE 42 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 11 OF 22 CAPLUS COPYRIGHT 2007 ACS on STN ACCESSION NUMBER: 1993:440374 CAPLUS Full-text

DOCUMENT NUMBER: 119:40374

TITLE: Inhibition of NF-KB activation by vitamin E

derivatives

AUTHOR(S): Suzuki, Yuichiro J.; Packer, Lester

CORPORATE SOURCE: Dep. Mol. Cell Biol., Univ. California, Berkeley, CA,

94720, USA

SOURCE: Biochemical and Biophysical Research Communications

(1993), 193(1), 277-83

CODEN: BBRCA9; ISSN: 0006-291X

DOCUMENT TYPE: Journal LANGUAGE: English

Nuclear factor KB (NF-KB) is believed to play an important role in the AB activation of a human immunodeficiency virus (HIV) which causes acquired immunodeficiency syndrome (AIDS). Recent findings suggesting an involvement of reactive oxygen species in signal transduction pathways leading to NF-KB activation have ensured the possible clin. use of antioxidants in blocking HIV activation. The present study examined the effects of vitamin E derivs. on the tumor necrosis factor- $\alpha$  (TNF- $\alpha$ )-induced NF- $\kappa B$  activation. Incubation of human Jurkat T cells with vitamin E acetate or  $\alpha\text{-tocopheryl}$  succinate (10  $\mu M$ to 1 mM) exhibited a concentration-dependent inhibition of NK-kB activation.  $\alpha\text{-Tocopherol}$  or succinate at these concns. had no apparent effects. 2,2,5,7,8-Pentamethyl-6-hydroxychromane (PMC) was extremely effective, causing complete inhibition of NK-kB activation at 10 µM. Oct-1 binding activity was inactivated by  $\alpha$ -tocopheryl succinate whereas other derivs. had no effects, suggesting that the effects of  $\alpha$ -tocopheryl succinate are not specific to NF- $\kappa B$ . HPLC measurements demonstrated that treatment of cells with TNF- $\alpha$  had no effects on cellular  $\alpha$ -tocopherol, but vitamin E acetate treatment increased the  $\alpha$ -tocopherol content. Cell viability was not affected by any of the vitamin E derivs. These results indicate a possible use of vitamin E derivs. in AIDS therapeutics.

IT 950-99-2

RL: BIOL (Biological study)

(TNF- $\alpha$ -induced nuclear factor  $\kappa B$  activation inhibition by,

AIDS therapy in relation to)

RN 950-99-2 CAPLUS

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

 $\begin{array}{c} \text{Me} \\ \text{Me} \\ \text{HO} \\ \text{Me} \end{array}$ 

L8 ANSWER 12 OF 22 USPATFULL on STN

ACCESSION NUMBER: 2006:289167 USPATFULL Full-text

TITLE: Use of active ingredients for the prophylaxis and/or

therapy of viral diseases

INVENTOR(S): Planz, Oliver, Wendelsheimer Strasse 34, Rottenburg,

GERMANY, FEDERAL REPUBLIC OF 72108

Pleschka, Stephan, Giessen, GERMANY, FEDERAL REPUBLIC

OF

Sedlacek, Hans-Harald, Marburg, GERMANY, FEDERAL

REPUBLIC OF

Ludwig, Stephan, Wurzeburg, GERMANY, FEDERAL REPUBLIC

OF

PATENT ASSIGNEE(S): Inamed GmbH Institut Fur Aerosolmedizin, Gemunden,

GERMANY, FEDERAL REPUBLIC OF, D-35285 (non-U.S.

corporation)

NUMBER DATE

PRIORITY INFORMATION: DE 2003-10300222 20030103

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: MAYER & WILLIAMS PC, 251 NORTH AVENUE WEST, 2ND FLOOR,

WESTFIELD, NJ, 07090, US

NUMBER OF CLAIMS: 15 EXEMPLARY CLAIM: 1 LINE COUNT: 781

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to the use preferably of at least one active ingredient for the prophylaxis and/or therapy of a viral disease, wherein this active ingredient inhibits at least one component of the cellular signal transduction pathway for the activation of the transcription factor NF-kB such that the virus multiplication is inhibited. The present invention relates furthermore to the local, preferably aerogenic administration of the active ingredient according to the invention for inhibiting a virus multiplication. The active ingredient according to the invention may be combined with at least one further antivirally effective substance for the prophylaxis and/or therapy of a viral disease.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2

(inhibitor of component of signal transduction for activation of NF-kB for prophylaxisnd/or therapy of virus diseases)

RN 950-99-2 USPATFULL

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

ANSWER 13 OF 22 USPATFULL on STN 1.8

2006:227433 USPATFULL Full-text ACCESSION NUMBER:

TITLE: Chroman derivatives as lipoxygenase inhibitors

Zhang, Wei, Santa Clara, CA, UNITED STATES INVENTOR(S):

Chen, Jian, San Jose, CA, UNITED STATES

Boddupalli, Sekhar, San Jose, CA, UNITED STATES

Galileo Pharmaceuticals, Inc (U.S. corporation) PATENT ASSIGNEE(S):

> NUMBER KIND DATE

PATENT INFORMATION: US 2006193797 A1 20060831

US 2006-349813 A1 20060207 (11) APPLICATION INFO.:

> NUMBER DATE \_\_\_\_\_

PRIORITY INFORMATION: US 2005-656644P 20050225 (60)

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

LEGAL REPRESENTATIVE: FOLEY & LARDNER LLP, 1530 PAGE MILL ROAD, PALO ALTO,

CA, 94304, US

NUMBER OF CLAIMS: 48 EXEMPLARY CLAIM: 1 2649 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention is concerned with certain novel derivatives of Formula AΒ wherein X and R.sup.1 to R.sup.10 are as described in the ##STR1## specification, and where either R.sup.5 is OH, --NR.sup.dOR.sup.a or --NR.sup.d--NR.sup.bR.sup.c, or R.sup.7 is --NR.sup.dOR.sup.a or --NR.sup.d--NR.sup.bR.sup.c, or C.dbd.R.sup.7R.sup.8 is C.dbd.NOR.sup.a or C.dbd.N--NR.sup.bR.sup.c, which may be useful in the manufacture of pharmaceutical compositions for treating disorders mediated by lipoxygenases. They may also be useful in the manufacture of skin care and/or pharmaceutical compositions for the treatment of lipoxygenase mediated disorders.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

950-99-2, 2,2,5,7,8-Pentamethylchroman-6-ol

(skin care and pharmaceutical compns. comprising chroman derivs. as lipoxygenase inhibitors)

RN 950-99-2 USPATFULL

2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME) CN

ANSWER 14 OF 22 USPATFULL on STN 1.8

2006:27860 USPATFULL Full-text ACCESSION NUMBER:

Novel pathways in the etiology of cancer TITLE:

Benz, Christopher C., Novato, CA, UNITED STATES INVENTOR(S): PATENT ASSIGNEE(S): Buck Institute for Age Research (U.S. corporation) NUMBER KIND DATE

PATENT INFORMATION: US 2006024691 A1 20060202

APPLICATION INFO.: US 2005-90546 Al 20050324 (11)

NUMBER DATE

PRIORITY INFORMATION: US 2004-556774P 20040325 (60)

US 2004-580534P 20040616 (60)

US 2004-629691P 20041119 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: QUINE INTELLECTUAL PROPERTY LAW GROUP, P.C., P O BOX

458, ALAMEDA, CA, 94501, US

NUMBER OF CLAIMS: 30 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 19 Drawing Page(s)

LINE COUNT: 2824

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention pertains to the identification of two novel epithelial signaling pathways in ER-positive breast cancer s and the discovery that the cellular biology and (likely also the clinical outcome) of ER-positive breast cancer cells is unexpectedly altered when these signaling pathways are activated. The first pathway pertains to the discovery that NF-KB activation and/or DNA binding is implicated in the etiology of ER-positive breast (and other) cancers. The second pathway involves ligand-independent quinine-mediated ER activation by posphorylation (e.g. on SER-118 and SER-167 residues of ER) and nuclear translocation of full-length (67 kDA) ER as well as the phorphorylating activation of a truncated and nuclear-localized ER variant (.about.52 kDa).

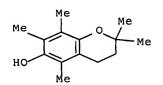
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2

(pathways in etiol. of cancer, and treatment methods)

RN 950-99-2 USPATFULL

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5;7,8-pentamethyl- (CA INDEX NAME)



L8 ANSWER 15 OF 22 USPATFULL on STN

ACCESSION NUMBER: 2005:221605 USPATFULL Full-text

TITLE: Chroman-derived anti-androgens for treatment of

androgen-mediated disorders

INVENTOR(S): Thompson, Todd A., Madison, WI, UNITED STATES

Wilding, George, Verona, WI, UNITED STATES

PATENT ASSIGNEE(S): Wisconsin Alumni Research Foundation (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 2005192342 A1 20050901

APPLICATION INFO.: US 2004-789835 A1 20040227 (10)

NUMBER DATE

PRIORITY INFORMATION: US 2003-450510P 20030227 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: GODFREY & KAHN, S.C., 780 N. WATER STREET, MILWAUKEE,

WI, 53202, US

NUMBER OF CLAIMS: 24 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 8 Drawing Page(s)

LINE COUNT: 1654

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Methods for the prevention and/or alleviation of androgen-mediated disorders treatable by administering a chroman-derived anti-androgen compound are provided by the present invention. The invention further provides pharmaceutical and nutraceutical compositions containing chroman-derived anti-androgen compounds useful in the prevention and/or alleviation of androgen-mediated disorders, particularly prostate cancer.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2, 2,2,5,7,8-Pentamethyl-6-chromanol

(chroman-derived antiandrogens for treatment of androgen-mediated disorders)

RN 950-99-2 USPATFULL

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

L8 ANSWER 16 OF 22 USPATFULL on STN

ACCESSION NUMBER: 2004:300069 USPATFULL Full-text

TITLE: Tocopherols, tocotrienols, other chroman and side chain

derivatives and uses thereof

INVENTOR(S): Sanders, Bob G., Austin, TX, UNITED STATES

Kline, Kimberly, Austin, TX, UNITED STATES Hurley, Laurence, Austin, TX, UNITED STATES Gardner, Robb, Austin, TX, UNITED STATES Menchaca, Marla, Austin, TX, UNITED STATES Yu, Weiping, Austin, TX, UNITED STATES

Ramanan, Puthucode N., Austin, TX, UNITED STATES

Liu, Shenquan, Austin, TX, UNITED STATES Israel, Karen, Austin, TX, UNITED STATES

PATENT ASSIGNEE(S): Research Development Foundation (U.S. corporation)

		NUMBER	KIND	DATE	
PATENT INFORMATION:	US	2004235938	Al	20041125	
APPLICATION INFO.:	US	2003-644418	A1	20030820	(10)

Division of Ser. No. US 2000-502592, filed on 11 Feb RELATED APPLN. INFO.:

2000, GRANTED, Pat. No. US 6770672 Continuation-in-part

of Ser. No. US 1999-404001, filed on 23 Sep 1999,

GRANTED, Pat. No. US 6417223

\_\_\_\_\_

NUMBER DATE

PRIORITY INFORMATION:

US 1998-101542P 19980923 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

Benjamin Aaron Adler, ADLER & ASSOCIATES, 8011 Candle

Lane, Houston, TX, 77071

NUMBER OF CLAIMS:

3.0

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

21 Drawing Page(s)

LINE COUNT:

2556

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides an antiproliferative compound having the AB

structural formula ##STR1##

wherein X is oxygen, nitrogen or sulfur; Y is selected from the group consisting of oxygen, nitrogen and sulfur wherein when Y is oxygen or nitrogen, n is 1 and when Y is sulfur, n is 0. Also provided is a method for inducing apoptosis in a cell comprising administering a composition comprising a compound.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2

(preparation of tocopherols, tocotrienols, other chroman and side chain derivs. for use as antitumor agents and for inducing cell apoptosis)

950-99-2 USPATFULL RN

2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME) CN

ANSWER 17 OF 22 USPATFULL on STN

ACCESSION NUMBER:

2004:192666 USPATFULL Full-text

TITLE:

Tocopherols, tocotrienols, other chroman and side chain

derivatives and uses thereof

INVENTOR(S):

Sanders, Bob G., Austin, TX, United States Kline, Kimberly, Austin, TX, United States Hurley, Laurence, Austin, TX, United States Gardner, Robb, Austin, TX, United States Menchaca, Marla, Austin, TX, United States Yu, Weiping, Austin, TX, United States

Ramanan, Puthucode N., Austin, TX, United States

Liu, Shenquan, Austin, TX, United States Israel, Karen, Austin, TX, United States

PATENT ASSIGNEE(S):

Research Development Foundation, Carson City, NV,

## United States (U.S. corporation)

APPLICATION INFO.: US 2000-502592 20000211 (9) RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1999-404001, filed on 23 Sep 1999, now patented, Pat. No. US 6417223,

issued on 9 Jul 2002

NUMBER DATE

PRIORITY INFORMATION: US 1998-101543P 19980923 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Fonda, Kathleen K. ASSISTANT EXAMINER: Maier, Leigh C.

LEGAL REPRESENTATIVE: Adler, Benjamin Aaron

NUMBER OF CLAIMS: 4 EXEMPLARY CLAIM: 1

PATENT INFORMATION:

NUMBER OF DRAWINGS: 18 Drawing Figure(s); 21 Drawing Page(s)

LINE COUNT: 2359

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention provides an antiproliferative compound having the structural formula ##STR1##

wherein X is oxygen, nitrogen or sulfur; Y is selected from the group consisting of oxygen, nitrogen and sulfur wherein when Y is oxygen or nitrogen, n is 1 and when Y is sulfur, n is 0. Also provided is a method for inducing apoptosis in a cell comprising administering a composition comprising a compound.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2, 2,2,5,7,8-Pentamethyl-6-chromanol

(preparation of tocopherols, tocotrienols, other chroman and side chain derivs. for therapeutic use in prevention and treatment of cancer)

RN 950-99-2 USPATFULL

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

L8 ANSWER 18 OF 22 USPATFULL on STN

ACCESSION NUMBER: 2004:127448 USPATFULL Full-text

TITLE: Tocopherols, tocotrienols, other chroman and side chain

derivatives and uses thereof

INVENTOR(S): Sanders, Bob G., Austin, TX, UNITED STATES

Kline, Kimberly, Austin, TX, UNITED STATES
Yu, Weiping, Austin, TX, UNITED STATES

PATENT ASSIGNEE(S): Research Development Foundation (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 2004097431 A1 20040520

APPLICATION INFO.: US 2003-695275 Al 20031028 (10)

RELATED APPLN. INFO.: Division of Ser. No. US 2001-8066, filed on 5 Nov 2001,

GRANTED, Pat. No. US 6703384 Continuation-in-part of Ser. No. US 2000-502592, filed on 11 Feb 2000, PENDING Continuation-in-part of Ser. No. US 1999-404001, filed

on 23 Sep 1999, GRANTED, Pat. No. US 6417223

NUMBER DATE

PRIORITY INFORMATION: US 1998-101542P 19980923 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Benjamin Aaron Adler, Ph.D., J.D., Adler & Associates,

8011 Candle Lane, Houston, TX, 77071

NUMBER OF CLAIMS: 17 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 11 Drawing Page(s)

LINE COUNT: 2605

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention provides an antiproliferative compound having a

structural formula ##STR1##

where X and Y independently are oxygen, nitrogen or sulfur; R.sup.1 is alkyl, alkenyl, alkynyl, aryl, heteroaryl, carboxylic acid, carboxylate, carboxamide, ester, thioamide, thiolacid, thiolester, saccharide, alkoxylinked saccharide, amine, sulfonate, sulfate, phosphate, alcohol, ethers or nitrites; R.sup.2 and R.sup.3 are hydrogen or R.sup.4; R.sup.4 is methyl, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzylester, saccharide or amine; and R.sup.5 is alkenyl; where when Y is nitrogen, said nitrogen is substituted with R.sup.6, wherein R.sup.6 is hydrogen or methyl. Also provided are methods for treating a cell proliferative disease and for inducing apoptosis in a cell comprising administering this compound is also provided.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2

(preparation of tocopherols, tocotrienols, other chroman and side chain derivs. for use as antitumor agents and for inducing cell apoptosis)

RN 950-99-2 USPATFULL

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

L8 ANSWER 19 OF 22 USPATFULL on STN

ACCESSION NUMBER: 2002:280579 USPATFULL Full-text

TITLE: Tocopherols, tocotrienols, other chroman and side chain

derivatives and uses thereof

INVENTOR(S): Sanders, Bob G., Austin, TX, UNITED STATES

Kline, Kimberly, Austin, TX, UNITED STATES Hurley, Laurence, Austin, TX, UNITED STATES Gardner, Robb, Austin, TX, UNITED STATES Menchaca, Marla, Austin, TX, UNITED STATES Yu, Weiping, Austin, TX, UNITED STATES

Ramanan, Puthucode N., Austin, TX, UNITED STATES

Liu, Shenquan, Austin, TX, UNITED STATES
Israel, Karen, Austin, TX, UNITED STATES

PATENT ASSIGNEE(S): Research Development Foundation (U.S. corporation)

US 6645998 B2 20031111

APPLICATION INFO.: US 2002-122019 A1 20020412 (10)

RELATED APPLN. INFO.: Division of Ser. No. US 1999-404001, filed on 23 Sep.

1999, GRANTED, Pat. No. US 6417223

NUMBER DATE

PRIORITY INFORMATION: US 1998-101542P 19980923 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Benjamin Aaron Adler, ADLER & ASSOCIATES, 8011 Candle

Lane, Houston, TX, 77071

NUMBER OF CLAIMS: 20 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 14 Drawing Page(s)

LINE COUNT: 2170

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention provides an antiproliferative compound having the

structural formula ##STR1##

wherein X is oxygen, nitrogen or sulfur; R.sup.1 is alkyl, alkenyl, alkynyl, aryl, heteroaryl, carboxylic acid, carboxylate, carboxamide, ester, thioamide, thiolacid, thiolester, saccharide, alkoxy-linked saccharide, amine, sulfonate, sulfate, phosphate, alcohol, ethers and nitriles; R.sup.2 is hydrogen, methyl, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzylester, saccharide and amine; R.sup.3 is selected from the group consisting of hydrogen, methyl, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzylester, saccharide and amine; R.sup.4 is of methyl, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzylester, saccharide and amine; and R.sup.5 is alkyl, alkenyl, alkynyl, aryl, heteroaryl, carboxyl, amide and ester. Also provided is a method for inducing apoptosis in a cell comprising administering a composition comprising a compound.

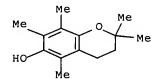
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2

RN

(preparation of tocopherols, tocotrienols, other chroman and side chain derivs. for use as antitumor agents and for inducing cell apoptosis) 950-99-2 USPATFULL

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)



ANSWER 20 OF 22 USPATFULL on STN L8

ACCESSION NUMBER: 2002:199098 USPATFULL Full-text

TITLE: Tocopherols, tocotrienols, other chroman and side chain

derivatives and uses thereof

Sanders, Bob G., Austin, TX, UNITED STATES INVENTOR(S):

Kline, Kimberly, Austin, TX, UNITED STATES

Yu, Weiping, Austin, TX, UNITED STATES

NUMBER KIND PATENT INFORMATION: US 2002107207 A1 20020808 B2 US 6703384 20040309 US 2001-8066 A1 20011105 (10) APPLICATION INFO.:

Continuation-in-part of Ser. No. US 2000-502592, filed RELATED APPLN. INFO.:

on 11 Feb 2000, PENDING Continuation-in-part of Ser. No. US 1999-404001, filed on 23 Sep 1999, PENDING

NUMBER DATE -----

PRIORITY INFORMATION: US 1998-101542P 19980923 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

Benjamin Aaron Adler, ADLER & ASSOCIATES, 8011 Candle LEGAL REPRESENTATIVE:

Lane, Houston, TX, 77071

NUMBER OF CLAIMS: 24 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 12 Drawing Page(s)

LINE COUNT: 2606

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention provides an antiproliferative compound having a

structural formula ##STR1##

where X and Y independently are oxygen, nitrogen or sulfur; R.sup.1 is alkyl, alkenyl, alkynyl, aryl, heteroaryl, carboxylic acid, carboxylate, carboxamide, ester, thioamide, thiolacid, thiolester, saccharide, alkoxylinked saccharide, amine, sulfonate, sulfate, phosphate, alcohol, ethers or nitrites; R.sup.2 and R.sup.3 are hydrogen or R.sup.4; R.sup.4 is methyl, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzylester, saccharide or amine; and R.sup.5 is alkenyl; where when Y is nitrogen, said nitrogen is substituted with R.sup.6, wherein R.sup.6 is hydrogen or methyl. Also provided are methods for treating a cell proliferative disease and for inducing apoptosis in a cell comprising administering this compound is also provided.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2, 2,2,5,7,8-Pentamethyl-6-chromanol

(preparation of tocopherols, tocotrienols, other chromans and side chain derivs. as potential antiproliferative, proapoptotic agents for the

RN 950-99-2 USPATFULL

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

L8 ANSWER 21 OF 22 USPATFULL on STN

ACCESSION NUMBER: 2002:168253 USPATFULL Full-text

TITLE: Tocopherols, tocotrienols, other chroman and side chain

derivatives and uses therof

INVENTOR(S): Sanders, Bob G., Austin, TX, United States

Kline, Kimberly, Austin, TX, United States Hurley, Laurence, Austin, TX, United States Gardner, Robb, Austin, TX, United States Menchaca, Marla, Austin, TX, United States

Yu, Weiping, Austin, TX, United States

Ramanan, Puthucode N., Austin, TX, United States

Liu, Shenquan, Austin, TX, United States Israel, Karen, Austin, TX, United States

PATENT ASSIGNEE(S): Research Development Foundation, Carson City, NV,

United States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 6417223 B1 20020709 APPLICATION INFO.: US 1999-404001 19990923 (9)

NUMBER DATE

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PRIORITY INFORMATION: US 1998-101542P 19980923 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Wilson, James O. ASSISTANT EXAMINER: Maier, Leigh C.

LEGAL REPRESENTATIVE: Adler, Benjamin Aaron

NUMBER OF CLAIMS: 3 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 14 Drawing Figure(s); 14 Drawing Page(s)

LINE COUNT: 1959

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention provides an antiproliferative compound having the

structural formula ##STR1##

wherein X is oxygen, nitrogen or sulfur; R.sup.1 is alkyl, alkenyl, alkynyl, aryl, heteroaryl, carboxylic acid, carboxylate, carboxamide, ester, thioamide, thiolacid, thiolester, saccharide, alkoxy-linked saccharide, amine, sulfonate, sulfate, phosphate, alcohol, ethers and nitriles; R.sup.2 is hydrogen, methyl, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzylester, saccharide and amine; R.sup.3 is selected from the group consisting of hydrogen, methyl, benzyl carboxylic acid, benzyl

carboxylate, benzyl carboxamide, benzylester, saccharide and amine; R.sup.4 is of methyl, benzyl carboxylic acid, benzyl carboxylate, benzyl carboxamide, benzylester, saccharide and amine; and R.sup.5 is alkyl, alkenyl, alkynyl, aryl, heteroaryl, carboxyl, amide and ester. Also provided is a method for inducing apoptosis in a cell comprising administering a composition comprising a compound.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2

(preparation of tocopherols, tocotrienols, other chroman and side chain derivs. for use as antitumor agents and for inducing cell apoptosis)

RN 950-99-2 USPATFULL

2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME) CN

ANSWER 22 OF 22 USPATFULL on STN

ACCESSION NUMBER:

92:55640 USPATFULL Full-text

TITLE:

Oxidized diphenylheteroalkanes

INVENTOR(S):

Janssen, Bernd, Ludwigshafen, Germany, Federal Republic

Wuest, Hans-Heiner, Dossenheim, Germany, Federal

Republic of

PATENT ASSIGNEE(S):

BASF Aktiengesellschaft, Ludwigshafen, Germany, Federal

Republic of (non-U.S. corporation)

NUMBER KIND DATE \_\_\_\_\_\_

PATENT INFORMATION:

US 5128479 19920707

APPLICATION INFO.:

US 1990-471886 19900129 (7)

NUMBER DATE -----

PRIORITY INFORMATION:

DE 1989-3903988 19890210

DOCUMENT TYPE: FILE SEGMENT:

Utility

Granted

PRIMARY EXAMINER:

Raymond, Richard L.

LEGAL REPRESENTATIVE: Oblon, Spivak, McClelland, Maier & Neustadt

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

1

1176

LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Oxidized diphenylheteroalkanes of the formula I ##STR1## where R.sup.1 to R.sup.6 and A have the meanings specified in the description, and the preparation thereof are described. The substances are suitable for

controlling diseases and as cosmetic agents.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 950-99-2P, 2,2,5,7,8-Pentamethylchroman-6-ol

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(preparation and reaction of, in preparation of drugs)
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RN 950-99-2 USPATFULL

CN 2H-1-Benzopyran-6-ol, 3,4-dihydro-2,2,5,7,8-pentamethyl- (CA INDEX NAME)

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(FILE 'HOME' ENTERED AT 12:23:49 ON 27 AUG 2007)

FILE 'REGISTRY' ENTERED AT 12:24:10 ON 27 AUG 2007

L1 STRUCTURE UPLOADED

L2 8 S L1 EXA FULL

FILE 'MEDLINE, CAPLUS, WPIDS, USPATFULL' ENTERED AT 12:24:41 ON 27 AUG 2007

L3 511 S L2

L4 11 S L3 AND "PROSTATE CANCER"

L5 1 S L3 AND "ANDROGEN-DEPENDENT"

FILE 'REGISTRY' ENTERED AT 12:43:27 ON 27 AUG 2007 E "PMCOL"/CN 25

FILE 'REGISTRY' ENTERED AT 12:44:06 ON 27 AUG 2007

L6 1 S 950-99-2/RN

SET NOTICE 1 DISPLAY
SET NOTICE LOGIN DISPLAY

FILE 'MEDLINE, CAPLUS, WPIDS, USPATFULL' ENTERED AT 12:44:38 ON 27 AUG 2007

L7 510 S L6

L8 22 S L7 AND (CANCER OR TUMOR)

---Logging off of STN---

=>
Executing the logoff script...

=> LOG Y

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COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
SINCE FILE TOTAL

ENTRY SESSION

CA SUBSCRIBER PRICE

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STN INTERNATIONAL LOGOFF AT 13:00:16 ON 27 AUG 2007